Type L# Hits				Search Text	DBs	Time Stamp mm Def ents init ion	Co or mm Def ents init ion	Err or Def init	Err
BRS L1 46050 antimicrobial		46050 antimicrobial	antimicrobial		USPAT; EPO; JPO; DERWENT	2003/09/13 10:58			To
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BRS L7 1 (5 or 6) and 2		1 (5 or 6) and 2	(5 or 6) and 2		USPAT; EPO; JPO; DERWENT	2003/09/13 11:02			0

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FILE 'AGRICOLA' ENTERED AT 11:05:24 ON 13 SEP 2003
=> s antimicrobial (p) (peptide or polypeptide)
           14203 ANTIMICROBIAL (P) (PEPTIDE OR POLYPEPTIDE)
L1
=> s polyphemusin-like
                2 POLYPHEMUSIN-LIKE
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DUPLICATE PREFERENCE IS 'CAPLUS, BIOSIS'
KEEP DUPLICATES FROM MORE THAN ONE FILE? Y/(N):n
PROCESSING COMPLETED FOR L2
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      ANSWER 1 OF 2 CAPLUS COPYRIGHT 2003 ACS on STN
13
                              2002:10505
ACCESSION NUMBER:
                                            CAPLUS
                              136:79729
DOCUMENT NUMBER:
                              Antimicrobial peptides and methods of use thereof
TITLE:
                              Hancock, Robert E. W.; Zhang, Lijuan
The University of British Columbia, Can.
INVENTOR(S):
PATENT ASSIGNEE(S):
                              PCT Int. Appl., 57 pp.
SOURCE:
                              CODEN: PIXXD2
DOCUMENT TYPE:
                              Patent
LANGUAGE:
                              English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                          KIND
                                                    APPLICATION NO.
      PATENT NO.
                                  DATE
                                                                         DATE
                                  20020103
      wo 2002000687
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                                                                         20010627
                                                    WO 2001-CA918
      wo 2002000687
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PRIORITY APPLN. INFO.:
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                                                                         20000627
                                                WO 2001-CA918
                                                                         20010627
OTHER SOURCE(S):
                              MARPAT 136:79729
      A class of cationic, ***polyphemusin*** - ***like*** peptides have antimicrobial activity is provided. Examples of such peptides include
                                                                              peptides having
      FRWCFRVCYKGRCRYKCR (SEQ ID NO:3), RRWCFRVCYKGFCRYKCR (SEQ ID NO:4), and RRWCFRVCYRGRFCYRKCR (SEQ ID NO:11) (I). Also provided are methods for inhibiting the growth of microbes such as bacteria, yeast and viruses utilizing the peptides of the invention. The peptides are particularly
      useful for inhibiting endotoxemia in a subject. I provided protection
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against endotoxemia in mice.

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COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN
                        BIOSIS
      ANSWER 2 OF 2
                        2002:159736 BIOSIS
ACCESSION NUMBER:
                        PREV200200159736
DOCUMENT NUMBER:
                        Antimicrobial peptides and methods of use thereof.
TITLE:
AUTHOR(S):
                        Hancock, Robert E. W. (1); Zhang, Lijuan
CORPORATE SOURCE:
                        (1) Vancouver Canada
                        ASSIGNEE: The University of British Columbia, Vancouver,
                        Canada
PATENT INFORMATION:
                       US 6337317 January 08, 2002
                        Official Gazette of the United States Patent and Trademark
SOURCE:
                        Office Patents, (Jan. 8, 2002) Vol. 1254, No. 2, pp. No.
                        Pagination. http://www.uspto.gov/web/menu/patdata.html.
                        ISSN: 0098-1133.
DOCUMENT TYPE:
                        Patent
LANGUAGE:
                        English
      A class of cationic,
                                 ***polyphemusin*** - ***like***
                                                                            peptides having
AB
      antimicrobial activity is provided. Examples of such peptides include
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      useful for inhibiting endotoxemia in a subject.
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=> s polyphemusin
             252 POLYPHEMUSIN
=> s hancock robert/au
               6 HANCOCK ROBERT/AU
=> s zhang lijuan/au
             201 ZHANG LIJUAN/AU
=> s 15 or 16
            207 L5 OR L6
=> s 17 and 14
               6 L7 AND L4
=> duplicate 18
ENTER REMOVE, IDENTIFY, ONLY, OR (?):remove DUPLICATE PREFERENCE IS 'CAPLUS, BIOSIS'
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KEEP DUPLICATES FROM MORE THAN ONE FILE? Y/(N):n
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PROCESSING COMPLETED FOR L8
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     ANSWER 1 OF 4
                       CAPLUS
                                 COPYRIGHT 2003 ACS on STN
                              2002:10505
ACCESSION NUMBER:
                                           CAPLUS
                              136:79729
DOCUMENT NUMBER:
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SOURCE:
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                              Patent
LANGUAGE:
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FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
      PATENT NO.
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                                 DATE
                                                 APPLICATION NO.
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20020103

Α2

WO 2001-CA918

20010627

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        wo 2002000687
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PRIORITY APPLN. INFO.:
                                                                WO 2001-CA918
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                                       MARPAT 136:79729
OTHER SOURCE(S):
       A class of cationic, ***polyphemusin*** -like peptides having antimicrobial activity is provided. Examples of such peptides include FRWCFRVCYKGRCRYKCR (SEQ ID NO:3), RRWCFRVCYKGFCRYKCR (SEQ ID NO:4), and RRWCFRVCYRGRFCYRKCR (SEQ ID NO:11) (I). Also provided are methods for including the growth of microbes such as bacteria, yeast and viruses included the invention. The poptides are particularly
        utilizing the peptides of the invention. The peptides are particularly useful for inhibiting endotoxemia in a subject. I provided protection
        against endotoxemia in mice.
                               BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN 2002:159736 BIOSIS PREV200200159736
        ANSWER 2 OF 4
ACCESSION NUMBER:
DOCUMENT NUMBER:
                                Antimicrobial peptides and methods of use thereof.
TITLE:
                                                                               ***Zhang, Lijuan***
                                Hancock, Robert E. W. (1);
AUTHOR(S):
                                (1) Vancouver Canada ASSIGNEE: The University of British Columbia, Vancouver,
CORPORATE SOURCE:
                                Canada
PATENT INFORMATION: US 6337317 January 08, 2002
SOURCE: Official Gazette of the United States Patent and Trademark
Office Patents, (Jan. 8, 2002) Vol. 1254, No. 2, pp. No
                                Pagination. http://www.uspto.gov/web/menu/patdata.html.
                                ISSN: 0098-1133.
DOCUMENT TYPE:
                                Patent
LANGUAGE:
                                English
                                            ***polyphemusin*** -like peptides having
        A class of cationic,
        antimicrobial activity is provided. Examples of such peptides include
        FRWCFRVCYKGRCRYKCR (SEQ ID NO:3), RRWCFRVCYKGFCRYKCR (SEQ ID NO:4), and RRWCFRVCYRGRFCYRKCR (SEQ ID NO:11). Also provided are methods for inhibiting the growth of microbes such as bacteria, yeast and viruses utilizing the peptides of the invention. The peptides are particularly
        useful for inhibiting endotoxemia in a subject.
        ANSWER 3 OF 4 CAPLUS
                                           COPYRIGHT 2003 ACS on STN DUPLICATE 1
ACCESSION NUMBER:
                                        2001:712236 CAPLUS
DOCUMENT NUMBER:
                                        136:49904
TITLE:
                                        Interaction of cationic antimicrobial peptides with
                                        model membranes
AUTHOR(S):
                                           ***Zhang, Lijuan*** ; Rozek, Annett; Hancock,
                                        Robert E. W.
                                        Department of Microbiology and Immunology, University of British Columbia, Vancouver, BC, V6T 1Z3, Can. Journal of Biological Chemistry (2001), 276(38),
CORPORATE SOURCE:
SOURCE:
                                        35714-35722
                                        CODEN: JBCHA3; ISSN: 0021-9258
PUBLISHER:
                                        American Society for Biochemistry and Molecular
                                        Biology
DOCUMENT TYPE:
                                        Journal
LANGUAGE:
                                        English
       A series of natural and synthetic cationic antimicrobial peptides from various structural classes, including .alpha.-helical, .beta.-sheet, extended, and cyclic, were examd. for their ability to interact with model
       membranes, assessing penetration of phospholipid monolayers and induction of lipid flip-flop, membrane leakiness, and peptide translocation across the bilayer of large unilamellar liposomes, at a range of peptide/lipid rare able to penetrate into monolayers made with neg. charged
       phospholipids, but only two interacted weakly with neutral lipids. Peptide-mediated lipid flip-flop generally occurred at peptide concns. that were 3- to 5-fold lower than those causing leakage of calcein across
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the membrane, regardless of peptide structure. With the exception of two .alpha.-helical peptides V681n and V25p, the extent of peptide-induced calcein release from large unilamellar liposomes was generally low at peptide/lipid molar ratios below 1:50. Peptide translocation across bilayers was found to be higher for the .beta.-sheet peptide ***polyphemusin*** , intermediate for .alpha.-helical peptides, and low for extended peptides. Overall, whereas all studied cationic antimicrobial peptides interacted with membranes, they were quite heterogeneous in their impact on these membranes.

RENCE COUNT: 35 THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
REFERENCE COUNT:
                                                                                                     RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
               ANSWER 4 OF 4 CAPLUS COPYRIGHT 2003 ACS ON STN DUPLICATE 2 SION NUMBER: 2000:769951 CAPLUS
ACCESSION NUMBER:
                                                                                 134:68617
DOCUMENT NUMBER:
                                                                                                                                       ***polyphemusin*** I and structural
                                                                                 Interaction of
TITLE:
                                                                                 analogs with bacterial membranes, lipopolysaccharide, and lipid monolayers

***Zhang, Lijuan***; Scott, Monisha G.; Yan, Hong
                                                                                                                                                              ; Scott, Monisha G.; Yan, Hong;
AUTHOR(S):
                                                                                 Mayer, Lawrence D.; Hancock, Robert E. W.
                                                                                 Department of Microbiology and Immunology, University of British Columbia, Vancouver, BC, V6T 1Z3, Can. Biochemistry (2000), 39(47), 14504-14514 CODEN: BICHAW; ISSN: 0006-2960
CORPORATE SOURCE:
SOURCE:
                                                                                 American Chemical Society
PUBLISHER:
                                                                                 Journal
DOCUMENT TYPE:
              UAGE: English

Three structural variants (PV5, PV7, and PV8) of the horseshoe crab cationic antimicrobial peptide ***polyphemusin*** I were designed with improved amphipathic profiles. CD spectroscopy anal. indicated that in phosphate buffer ***polyphemusin*** I, PV7, and PV8 displayed the spectrum of a type II .beta.-turn-rich structure, but, like

***polyphemusin*** I, all three variants adopted a typical .beta.-sheet structure in an anionic lipid environment. Both ***polyphemusin*** I and variants were potent broad spectrum antimicrobials that were clearly bactericidal at their minimal inhibitory concns. The variants were moderately less active in vitro but more effective in animal models.

Moreover, these variants exhibited delayed bacterial killing, whereas

***polyphemusin*** I killed Escherichia coli UB1005 within 5 min at 2.5
.mu.g/mL. All the peptides showed similar abilities to bind to bacterial lipopolysaccharide (LPS) and permeabilize bacterial outer membranes.

Consistent with this was the observation that all peptides significantly inhibited cytokine prodn. by LPS-stimulated macrophages and penetrated polyanionic LPS monolayers to similar extents. None of the peptides had affinity for neutral lipids as evident from both tryptophan fluorescence
LANGUAGE:
                                                                                 English
                affinity for neutral lipids as evident from both tryptophan fluorescence
               spectroscopy and Langmuir monolayer anal. As compared to

***polyphemusin*** I, all variants showed reduced ability to interact
with anionic lipids, and the hemolytic activity of the variants was
decreased by 2-4-fold. In contrast, ***polyphemusin*** I efficiently
               decreased by 2-4-fold. In contrast, ***polyphemusin*** I efficiently depolarized the cytoplasmic membrane of Escherichia coli, as assessed using a membrane potential sensitive fluorescent dye 3,3-dipropylthiacarbocyanine (disC35) assay, but the variants showed a substantially delayed and decreased depolarizing ability. The coincident assessment of cell viability indicated that depolarization of the bacterial cytoplasmic membrane potential by ***polyphemusin*** I occurred prior to lethal damage to cells. Our data suggest that increase of amphipathicity of .beta.-sheet ***polyphemusin*** I generally resulted in variants with decreased activity for membranes.
                Interestingly, all variants showed an improved ability to protect mice both against infection by Pseudomonas aeruginosa and from endotoxemia. ENCE COUNT:

41 THERE ARE 41 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                                                                                                     RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
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               FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH, AGRICOLA' ENTERED AT 11:05:24 ON 13 SEP 2003
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                                                   S POLYPHEMUSIN-LIKE
                                              2 DUPLICATE REMOVE L2 (0 DUPLICATES REMOVED)
                                      252 S POLYPHEMUSIN
                                            6 S HANCOCK ROBERT/AU
                                      201 S ZHANG LIJUAN/AU
207 S L5 OR L6
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6 S L7 AND L4

L8 L9